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1101 CCTTTGACTC CTCGGAGCC CTCATGAGGA AGTTGGGCT CATGGACAAAT GAGATAAGG TGGCTAAAGC TGAGGAGGG GCCCACAGGG ACACCTGTA
 322 GAAACTGAG GACCTGGG GACTACTCTC TCAACCCGGA GTACGTTGTTA CCTTATTCG ACGGAATTTGG ACTOCGCTGCC CCGGTGTCCTC TGTGAAACAT
 322 PheAsp 85 FEPGLUPRO LeuMetArg YsLeuGlyLeu MetAspSer GluIleLysV AlAIAlySAlAgluAlaLia GlyHisArgA AsPheLeutYr
 1201 CACCATGCTG ATAAAGTGGG TCAACAAAC CGGGCGGAGAT GCCTCTGCTC ACACCCGCTGT GGATGCGCTTG GAGACGCTGG GAGAGAGACT TGCCAAGCAG
 355 GCGCTAGAC TATTCAACCC AGTTGTTTC GCGCGCTCTA CGGAGACAGG TGTTGGACCA CCTACGGAAC CCTCTGGACCC CTCTCTCTGA ACGGTTGTC
 ThemetLiu IleIlyStpY alAsnLysThr ArgIleArgAsp AlaservAlh IleThrLeuL LeuIleAlaL GluThrLeuL GluIleArgLeuL ValAlaLysGln
 1301 AGGATTGAGG ACCACTTGTG GAGCTCTGG AAGTTCATGT ATCTAGAAGG TAATGAGG TCTGCGCTGTG ATTCAGCTTCA GGAAGTGGAGA
 TCTTAACTC CTTGTAACAA CTCGAGACCT TTCAAGTACA TAGATCTTCC ATTACGTCCTG AGACGGAAC TAAGAGAAGT CCTTCACCTCT
 388 LysIleGlu SphIleLeu userserGly LysPheMetT YIleUglI YIleUglI YIleUglI YIleUglI YIleUglI YIleUglI YIleUglI YIleUglI
 1401 CCTTCCCTGC TTACTCTTT TTCTGGAAA AGCCCAACTG GACTCCAGTC AGTAGGAAAG TCCCACAAATT GTCACATGAC CGGTACTGGAA AGAAACTCTC
 GAAAGGACC AAATGGAAA AGACACTTT TCGGGTGAC CTGAGGTGAG TCATCCTTTC ACGGTTAA CAGTGTACTG GCCATGACCT TCTTTGAGAG
 1501 CCATCCAACA TCACCCAGTG GATGGAACAT CCTGTAACCTT TTCACTGTCAC TTGGCATTAT TTATATAAGC TGAAATGTCAT AATAAGGACA CTATGGAAAT
 CGTAGGGTTT AGTGGTCAC CTACCTGTAA GGACATTGAA AACGCTTAATA AACCGTAAACTA AACGCTTAACACTA TTATTCCTGT GATAACCTTTA
 1601 CTCTGGATCA TTCCGGTTGT GCGTACTTTG AGATTGGTT TGGGATGTCAC TTGTTTTTCAC AGCACCTTTT TATCCTAAATG TAATATGCTTT ATTTATTTAT
 CAGACCTAGT AAGGCAAAAC CCCATGAAAC TCTAAACCA ACCCTACAGT AACAAAAAGTG TCGTGAAGAATAGGATTAC ATAGGATTAC ATTTAGAAA TAAATAATA
 1701 TTGGGCTACA TTGTAAGATC CATCTACAAA AAAAAGGAAAGG AAAAAGGAAAGG AAAAAGGAAAGG AAAAAGGAAAGG AAAAAGGAAAGG
 AACCCCGATGG AACATTCCTAG GTAGATGTTT TTPTTTTTTTT TTPTTTTTTTT TTPTTTTTTTT TTPTTTTTTTT TTPTTTTTTTT TTPTTTTTTTT
 CGACCTGTCAC CGGGGGGGGG ACTCTAGAGT CGACCTGGCAG AAGCTGGGCC GGCATGGCC GCTGGACGTC TTGAGATCTCA GTCAGTACCGG
 CGGTACCGG

Fig. 1 (cont.)

Fig 2A

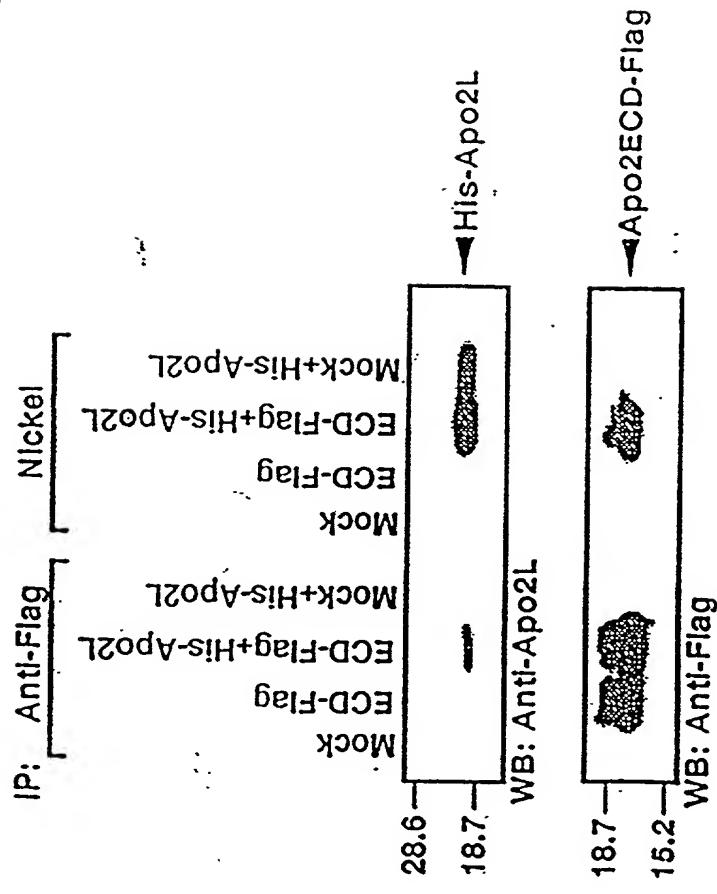
1 MEORGONAPAAASGARKRHHGPGPREARGARPGLRVPKTLVLLVVAEPLLVAESALITQQD
 61 LAPQQRAAPQQKRSPPSEGICPPGHHSIEDGRDCISCKYQDYSSTHWNDLFLFQLRCTRQD
 121 SGEVELSPCTTRNTVCOCEEGTFREEDSPEMCRKCRITGCPRGMVKGPRGMSQRRLLVPA
 181 KESGIIIGTVAAVVLIVAVEVCKSLIWKVLPYLGICSGGGDPERVDRSSQRPGaed
 241 NVLINEITVSILQOPTQVPEQEMEVQEPAAEPTGVNMLSPGESEHLLPEAEERSQRRLVPA
 301 NEGDPTETRQCFDDFAADLVFDSWEPLMRKLGMDNEIKYAKAAEAGHRDTLYTMLIKW
 361 VNKTRGRDASVHTLLDAETLGERLAKOKIEDHLLSSGKFMYLEGNADSALS

Fig 2B

Apo2 DR4 Apo3/DR3 TNFR1 Fas/Apo1
 EADL VPEPDKDQLMRQIDITKNEVTKTGRKREAEVTKTGRVNEAKIDEDIKNDN
 FNI VMDA VVEN IGVMTLSQVNG
 VPEPDKDQLMRQIDITKNEVTKTGRKREAEVTKTGRVNEAKIDEDIKNDN
 VMDA VVEN IGVMTLSQVNG
 VPEPDKDQLMRQIDITKNEVTKTGRKREAEVTKTGRVNEAKIDEDIKNDN
 VMDA VVEN IGVMTLSQVNG

Apo2 DR4 Apo3/DR3 TNFR1 Fas/Apo1
 VTIKRNTRGTD - ASVHDTDATEBLAKQKIED
 VAMKRNTRGDN - ASIHLIDDALEERHAKQKIED
 VELKRNTRGQP - AGTGAIVYAALEERHAKQKIED
 VCATNRRTRPPEATI - GELGRVFRDMDFLGCLQKIED
 - QLNRNHHQLHCKKEAY - DTHIKDEKKANEKLTAEKQKIED

Fig. 3



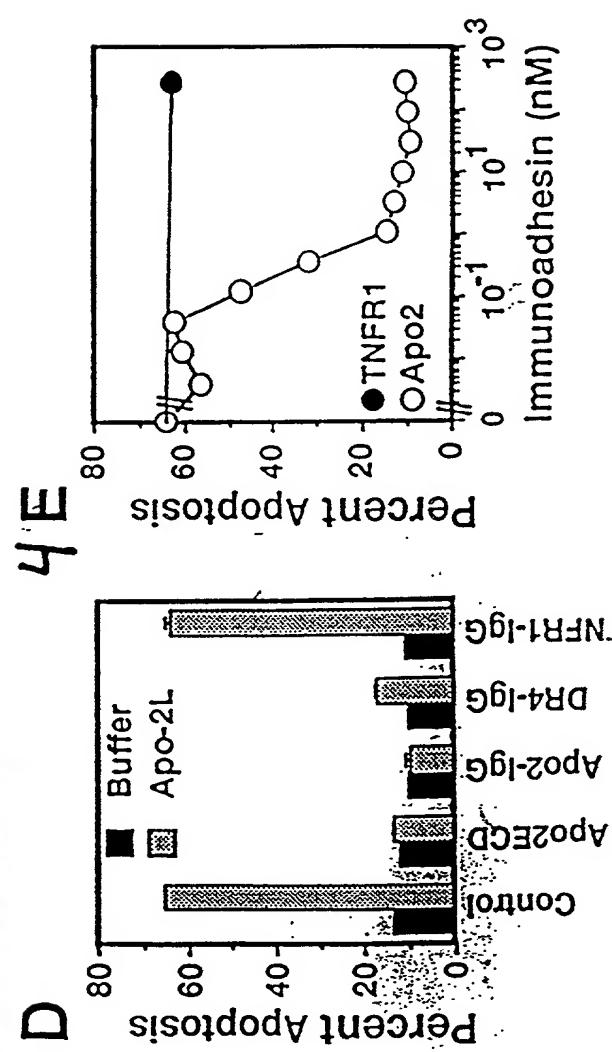
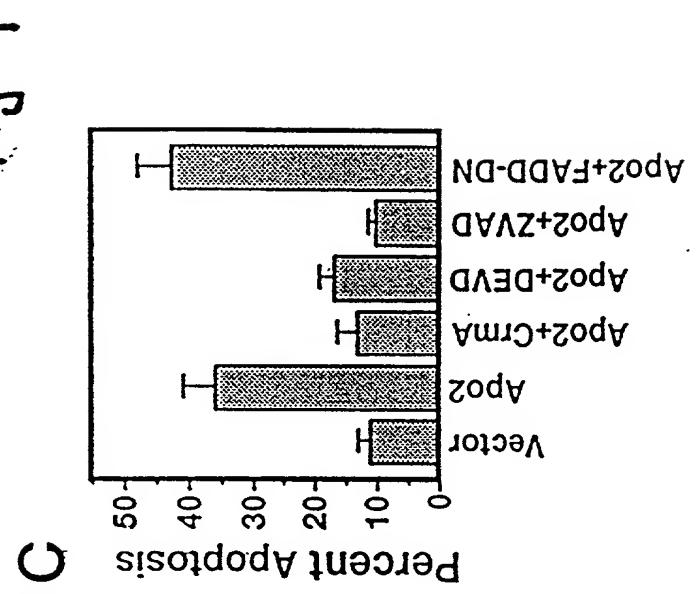
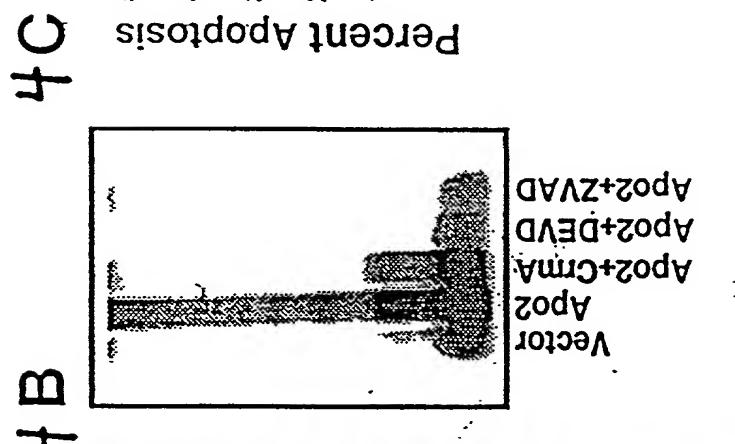
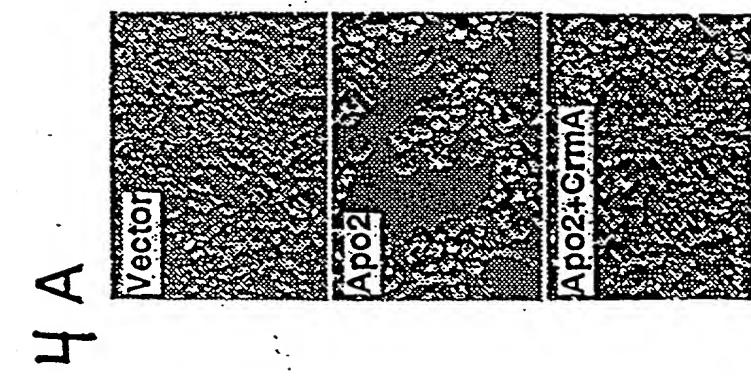


Fig. 4

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5C
5B

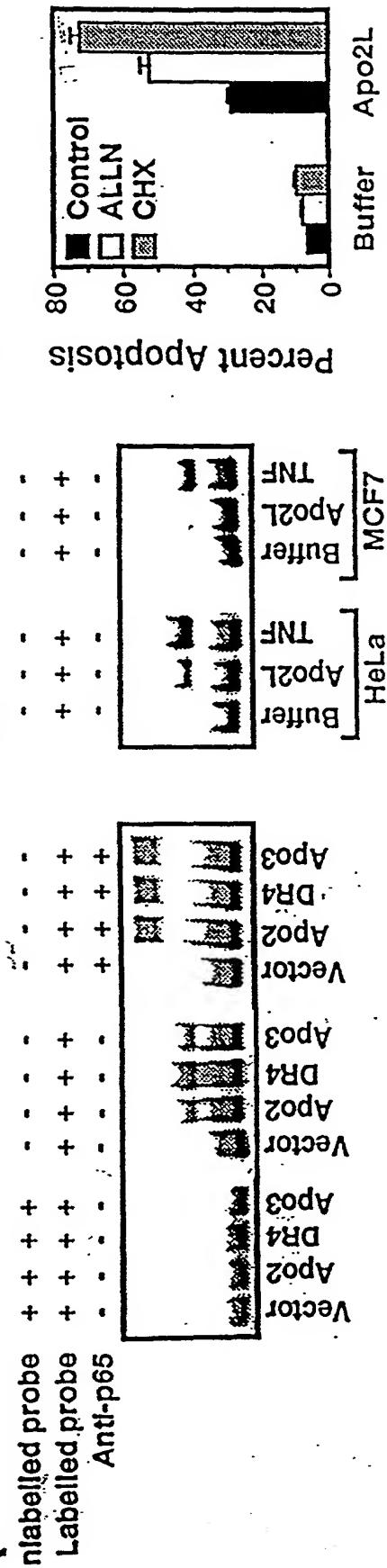
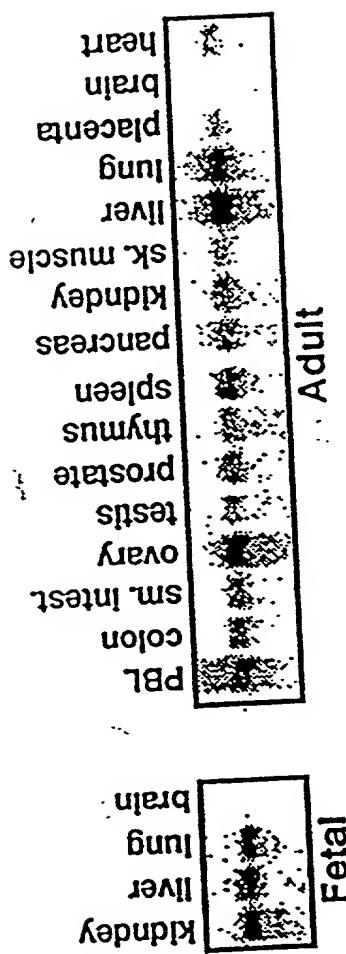
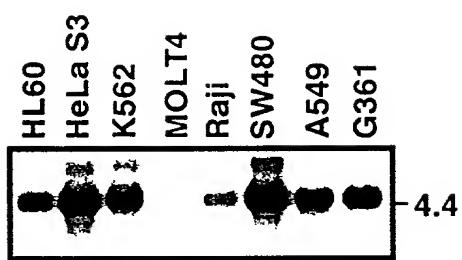


Fig. 5

FIG. 6A





Cancer cell line

Fig. 6B

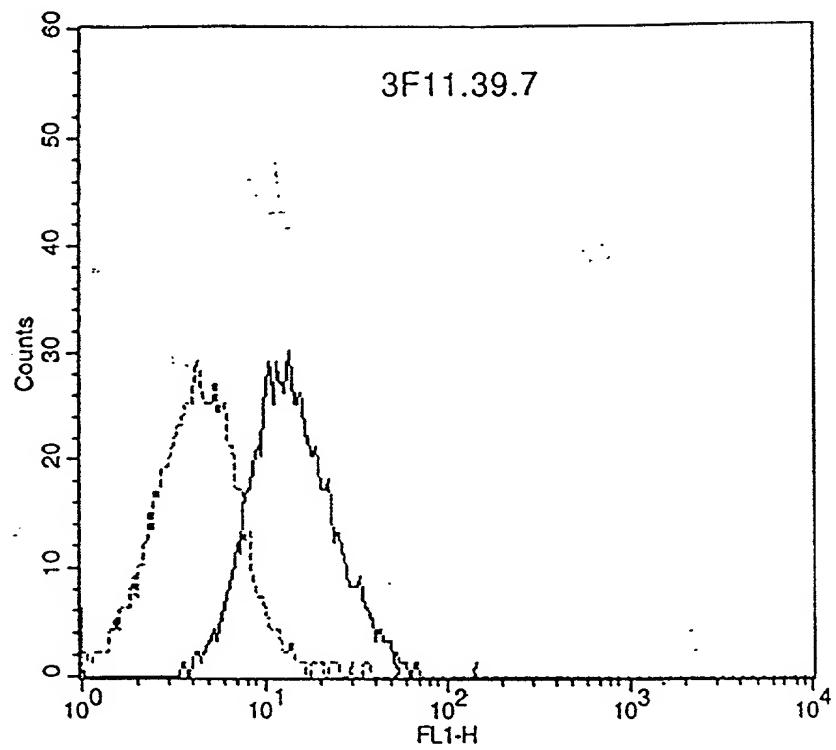


Fig. 7

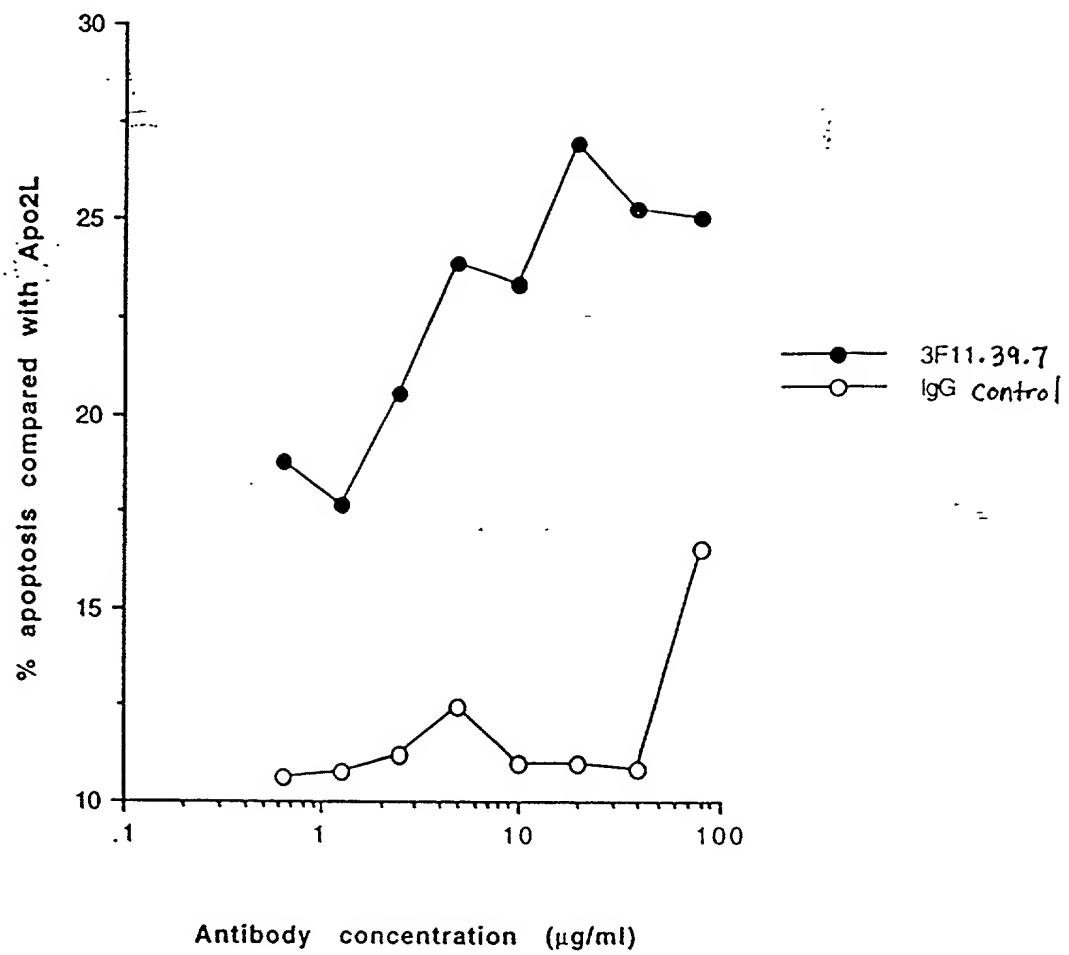


Fig. 8

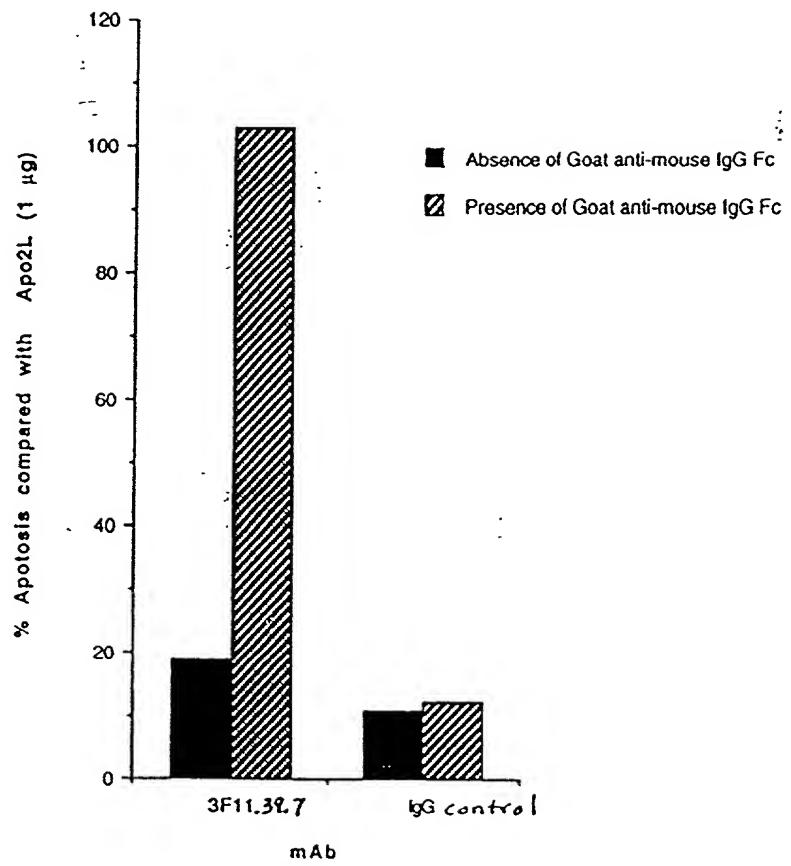


Fig. 9

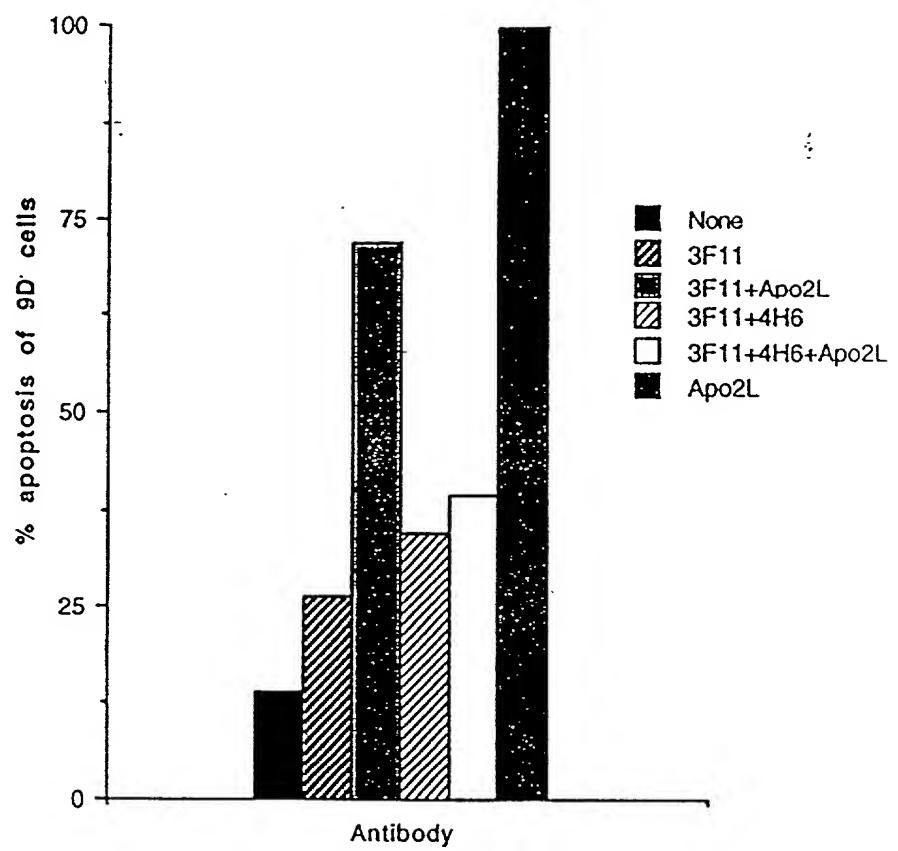


Fig : 10

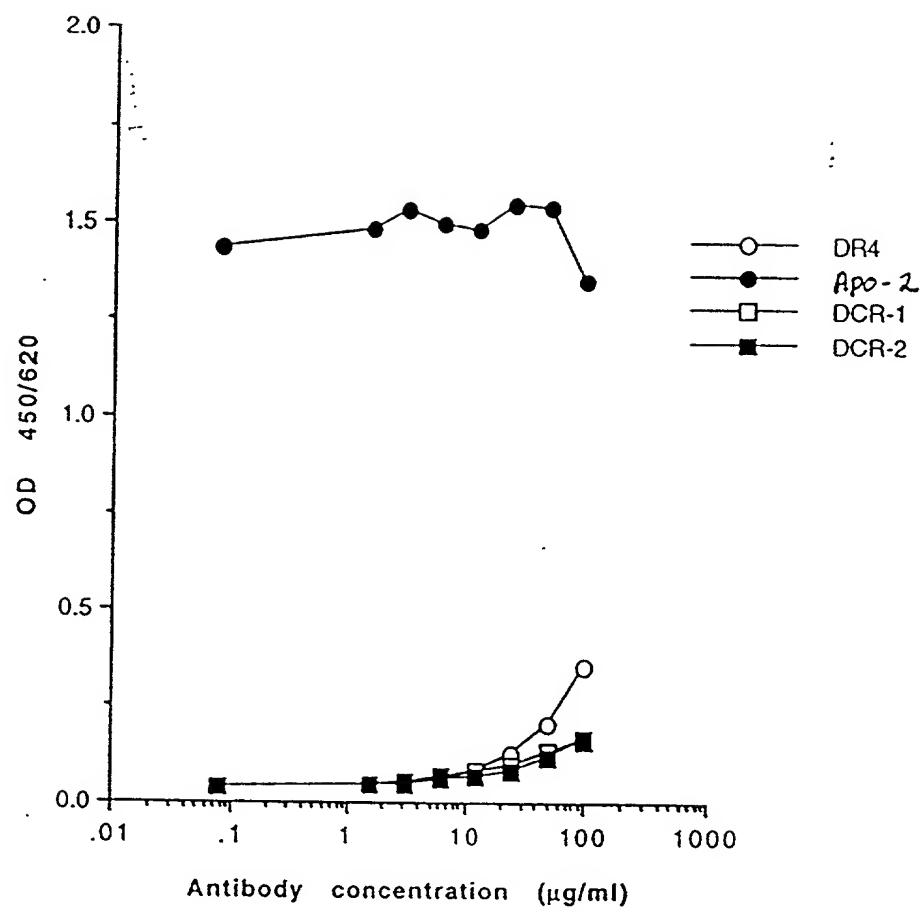


Fig. 11

anti-Apo-2 16E2-his

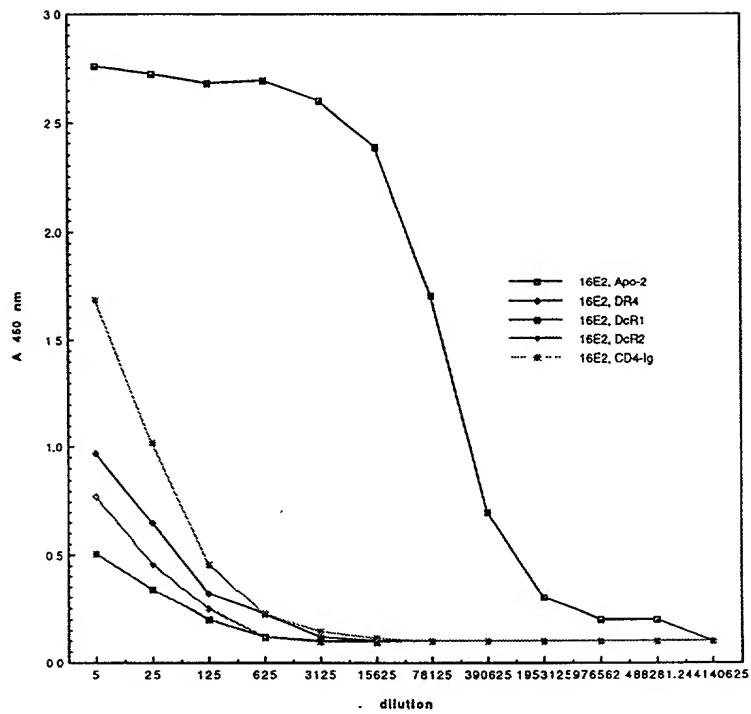


Fig. 12A

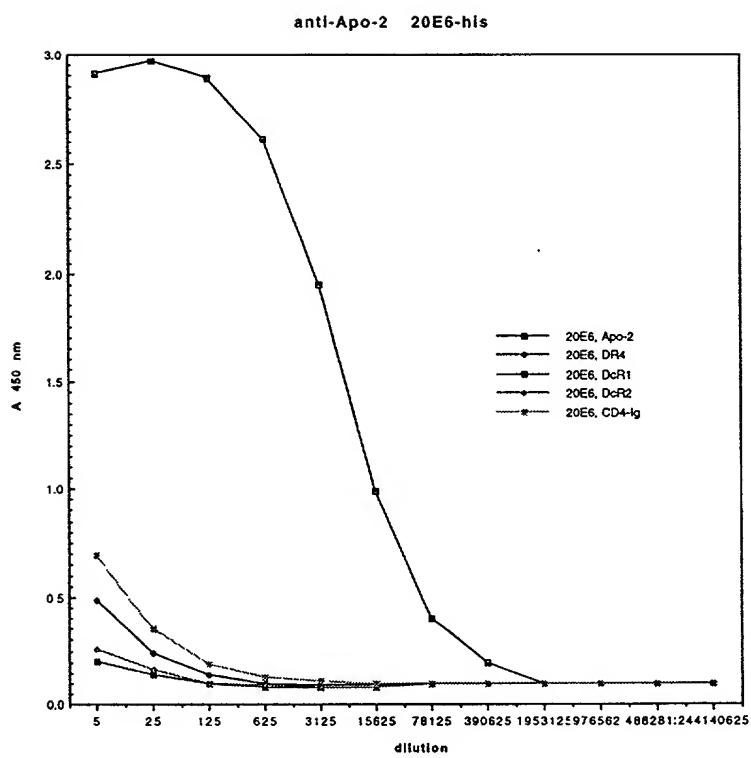


Fig. 12B

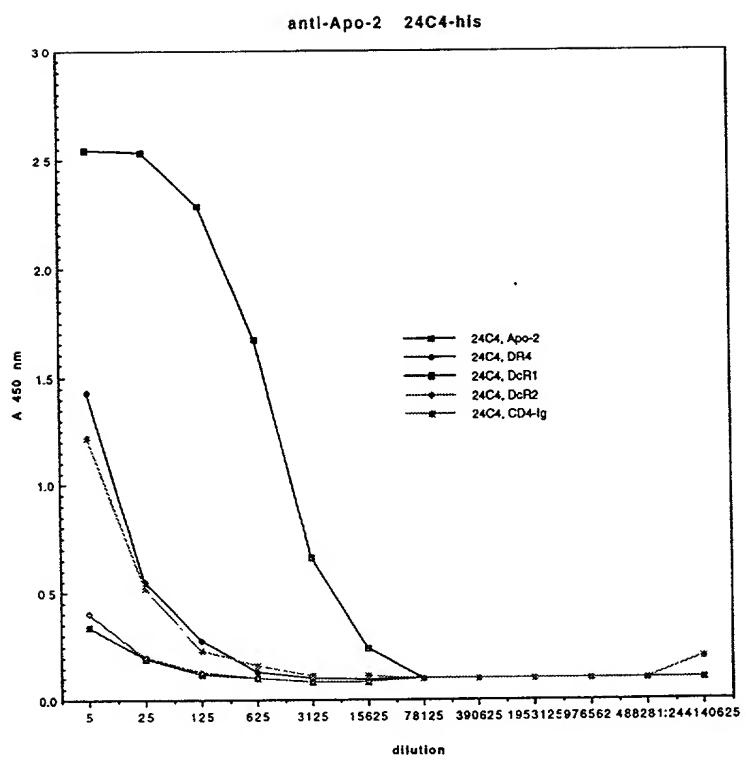


Fig. 12C

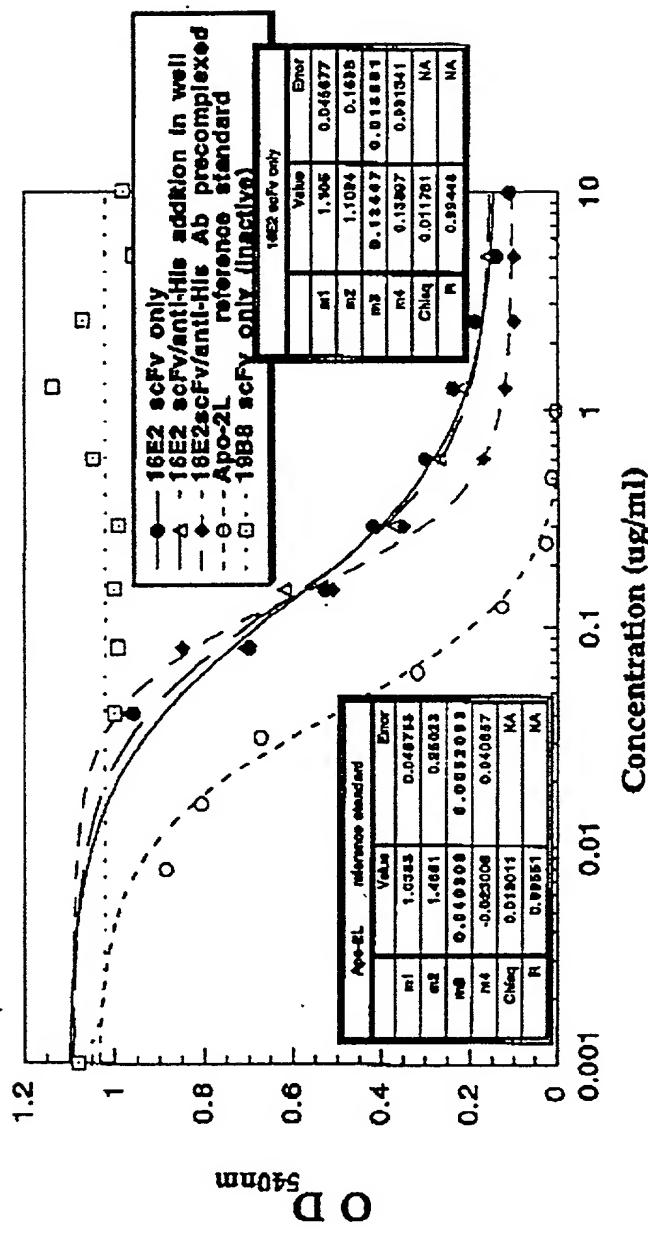


Fig. 13A

Fig. 13B

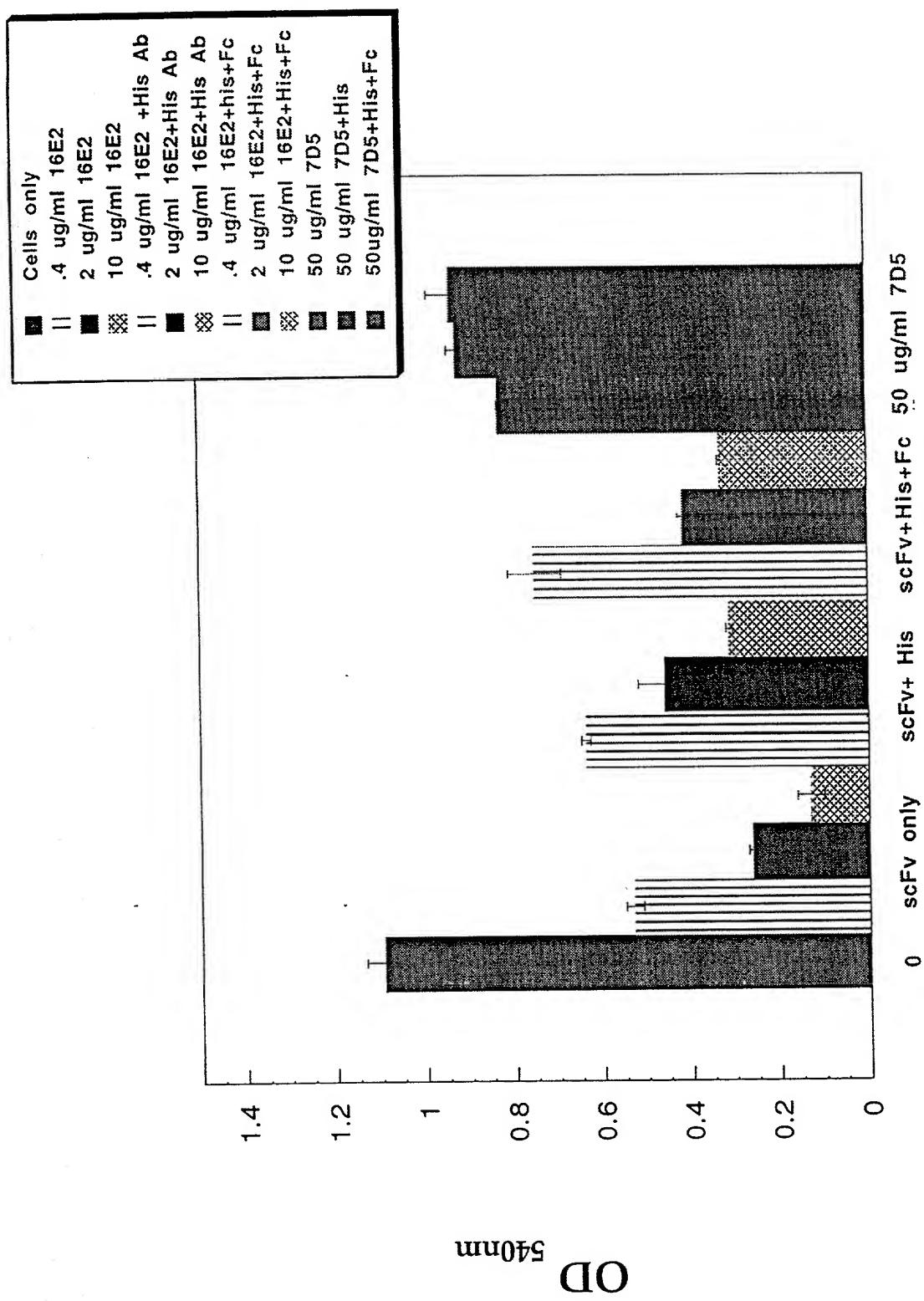
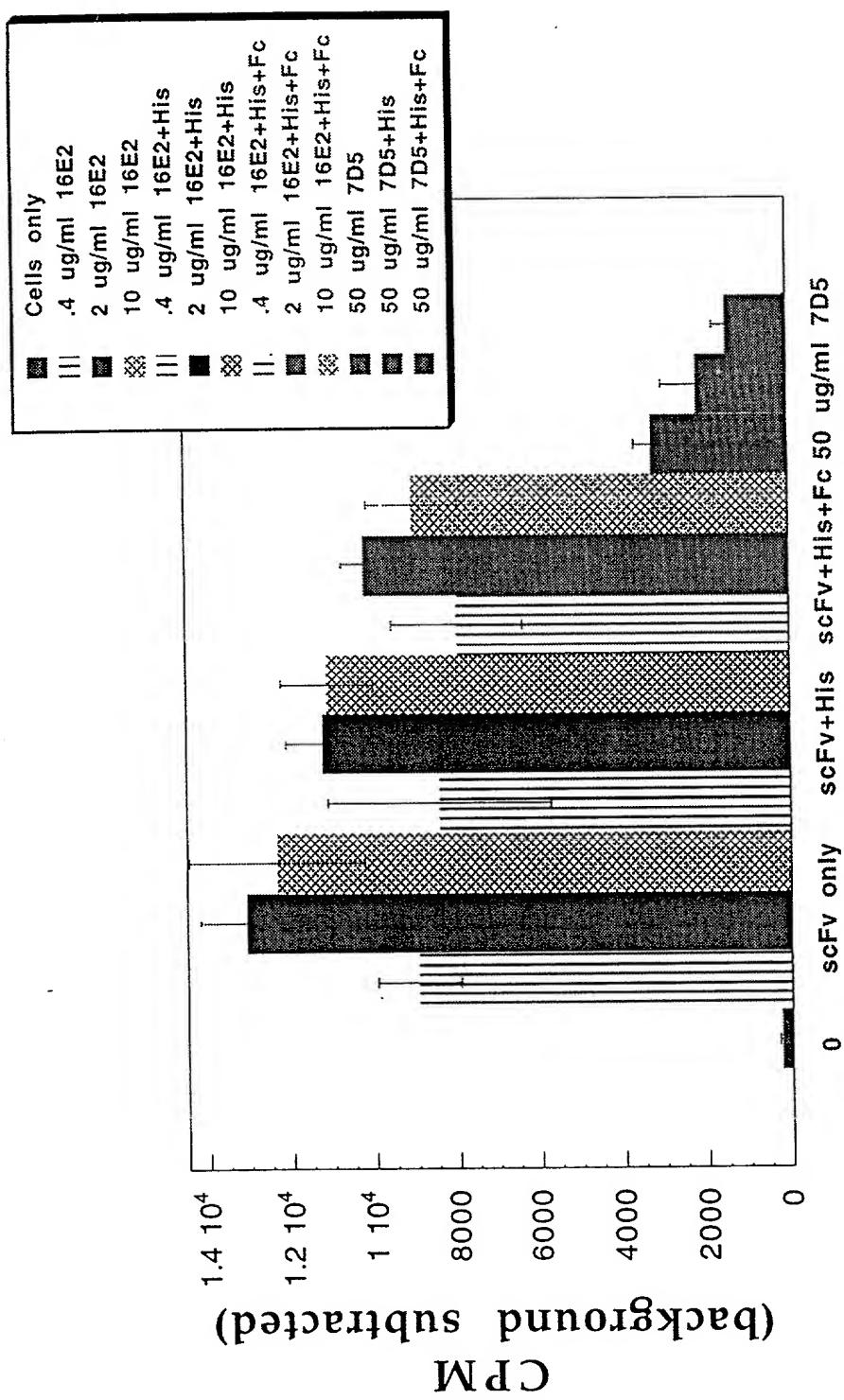


Fig. 13c



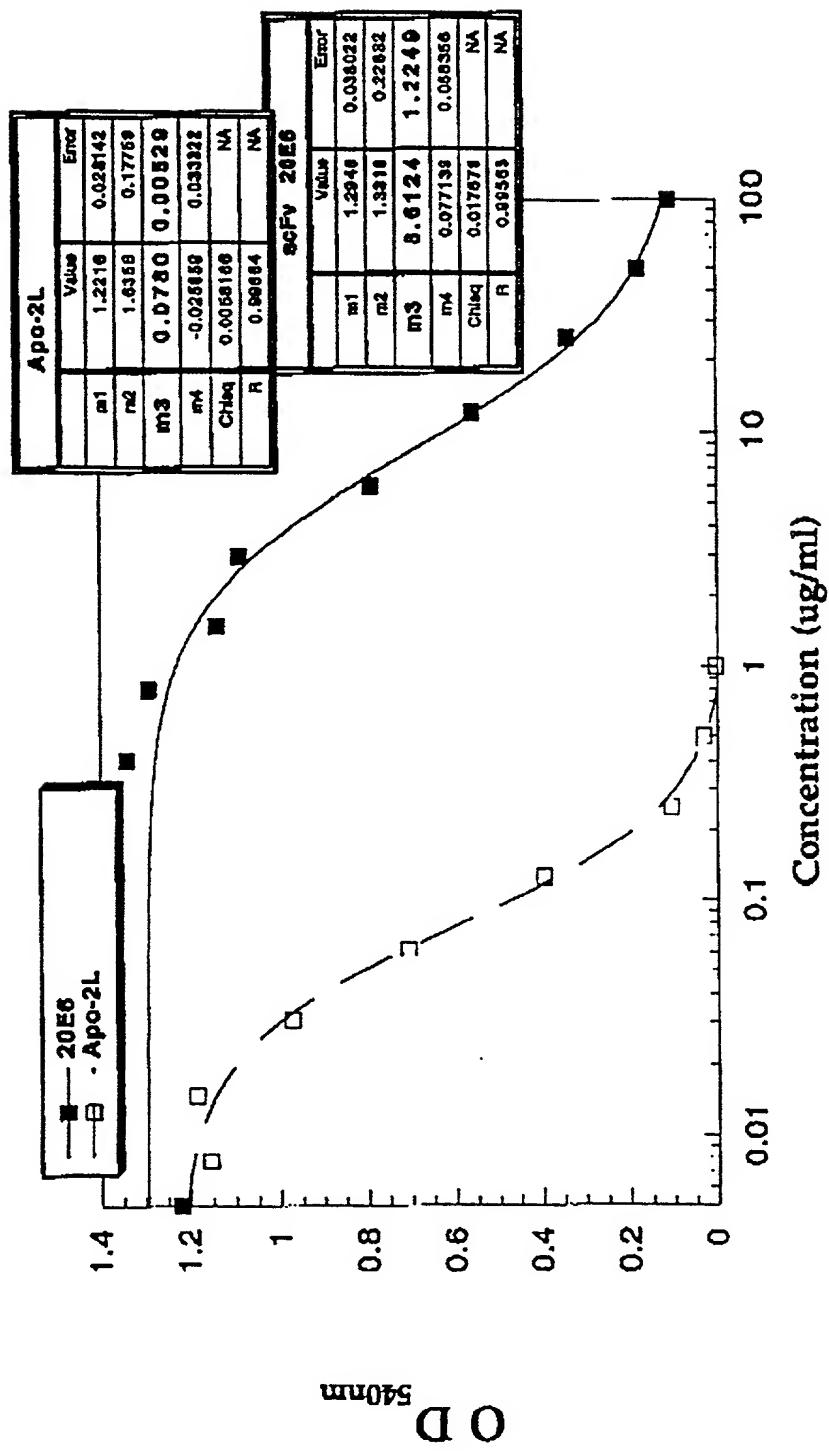


Fig 14A

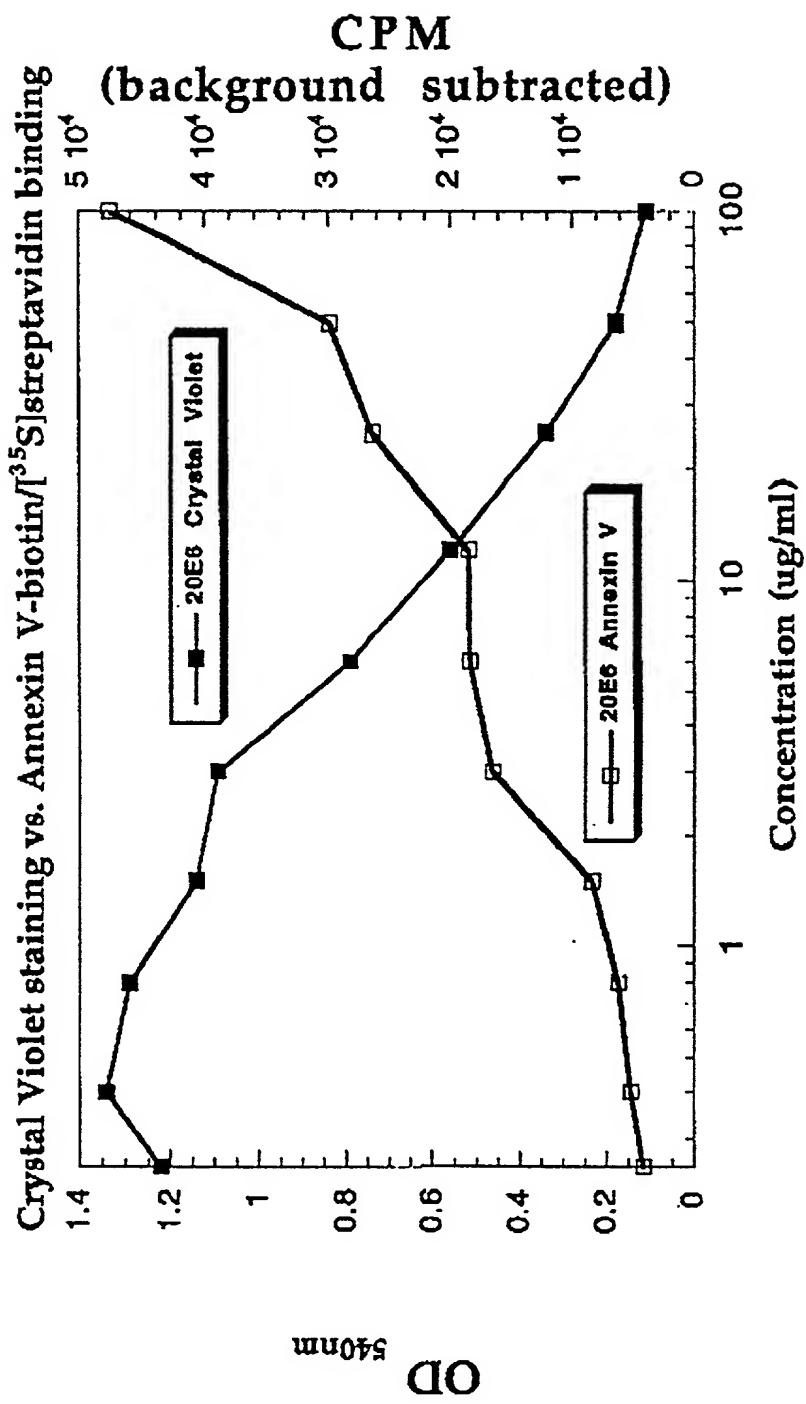
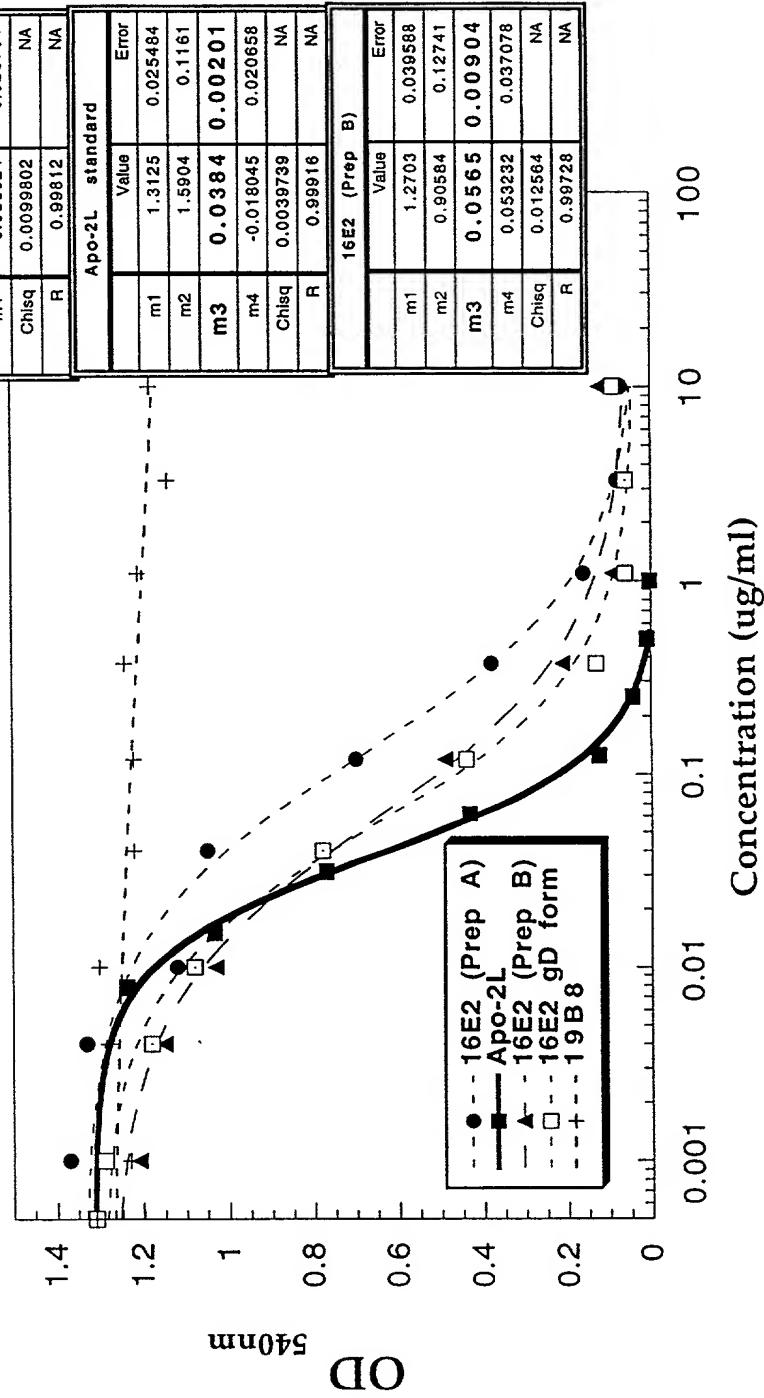


Fig. 148

Fig. 14C



ATGACCATGA TTACGCCAAG CTTTGGAGCC TTTTTTTGG AGATTTCAA 50
CGTGAAAAAA TTATTATTCTG CAATTCTTT AGTTGTCCT TTCTATGCGG 100
CCCAGCCGGC CATGGCCGAG GTGCAGCTGG TGCAGTCTGG GGGAGGTGTG 150
GAACGGCCGG GGGGGTCCCT GAGACTCTCC TGTGCAGCCT CTGGATTAC 200
CTTTGATGAT TATGGCATGA GCTGGTCCG CCAAGCTCCA GGGAAAGGGC 250
TGGAGTGGGT CTCTGGTATT AATTGGAATG GTGGTAGCAC AGGATATGCA 300
GAATCTGTGA AGGGCCGAGT CACCCTCTCC AGAGACAACG CCAAGAACTC 350
CCTGTATCTG CAAATGAACA GCCTGAGAGC CGAGGACACG GCCGTATATT 400
ACTGTGCGAA AATCCTGGT GCCGGACGGG GCTGGTACTT CGATCTCTGG 450
GGGAAGGGGA CCACGGTCAC CGTCTCGAGT GGTGGAGGCG GTTCAGGCGG 500
AGGTGGCAGC GGCGGTGGCG GATCGTCTGA GCTGACTCAG GACCCTGCTG 550
TGTCTGTGGC CTTGGGACAG ACAGTCAGGA TCACATGCCA AGGAGACAGC 600
CTCAGAAGCT ATTATGCAAG CTGGTACCAAG CAGAAGCCAG GACAGGCC 650
TGTACTTGTC ATCTATGGTA AAAACAACCG GCCCTCAGGG ATCCCAGACC 700
GATTCTCTGG CTCCAGCTCA GGAAACACAG CTTCCCTGAC CATCACTGGG 750
GCTCAGGCGG AAGATGAGGC TGACTATTAC TGTAACCTCCC GGGACAGCAG 800
TGGTAACCAT GTGGTATTCTG GCGGAGGGAC CAAGCTGACC GTCCTAGGTG 850
CGGCCGCACA TCATCATCAC CATCACGGGG CCGCAGAACAA AAAACTCATC 900
TCAGAAGAGGG ATCTGAATGG GGCCGCATAG 930

Fig. 15A

40952736 210202

ATGACCATGA TTACGCCAAG CTTTGGAGCC TTTTTTTGG AGATTTCAA 50
CGTGAAAAAA TTATTATTCTG CAATTCTTT AGTTGTCCT TTCTATGCGG 100
CCCAGCCGGC CATGGCCGGG GTGCAGCTGG TGGAGTCTGG GGGAGGCTTG 150
GTCCAGCCTG GGGGGTCCCT GAGACTCTCC TGTGCAGCCT CTGGATTAC 200
CTTTAGTAGC TATTGGATGA GCTGGGTCCG CCAGGCTCCA GGGAAAGGGC 250
TGGAGTGGGT GGCCAACATA AAGCAAGATG GAAGTGAGAA ATACTATGTG 300
GAECTCTGTGA AGGGCCGATT CACCCTCTCC AGAGACAACG CCAAGAACTC 350
ACTGTATCTG CAAATGAACA GCCTGAGAGC CGAGGACACG GCTGTGTATT 400
ACTGTGCGAG AGATCTTTA AAGGTCAAGG GCAGCTCGTC TGGGTGGTTC 450
GACCCCTGGG GGAGAGGGAC CACGGTCACC GTCTCGAGTG GTGGAGGCGG 500
TTCAGGCGGA GGTGGTAGCG GCGGTGGCGG ATCGTCTGAG CTGACTCAGG 550
ACCCTGCTGT GTCTGTGGCC TTGGGACAGA CAGTCAGGAT CACATGCCAA 600
GGAGACAGCC TCAGAAGCTA TTATGCAAGC TGGTACCAAGC AGAAGCCAGG 650
ACAGGCCCT GTACTTGTCA TCTATGGTAA AAACAACCGG CCCTCAGGGA 700
TCCCAGACCG ATTCTCTGGC TCCAGCTCAG GAAACACAGC TTCCTTGACC 750
ATCACTGGGG CTCAGGCGGA AGATGAGGCT GACTATTACT GTAACTCCCG 800
GGACAGCAGT GGTAACCATG TGGTATTCTGG CGGAGGGACC AAGCTGACCG 850
TCCTAGGTGC GGCGCACAT CATCATCACC ATCACGGGGC CGCAGAACAA 900
AAACTCATCT CAGAAGAGGA TCTGAATGGG GCCGCATAG 939

Fig. 15B

ATGACCATGA TTACGCCAAG CTTTGGAGCC TTTTTTTGG AGATTTCAA 50
CGTGAAAAAA TTATTATTAG CAATTCTTT AGTTGTTCCCT TTCTATGCAGG 100
CCCAGCCGGC CATGGCCAG GTGCAGCTGG TGCACTCTGG GGGAGGCCTG 150
GTCCAGCCTG GGCGGTCCCT GAGACTCTCC TGTGCAGCTT CTGGGTTCAT 200
TTTCAGTAGT TATGGGATGC ACTGGGTCCG CCAGGCTCCA GGCAAGGGC 250
TGGAGTGGGT GGCAAGGTATT TTTTATGATG GAGGTAATAA ATACTATGCA 300
GACTCCGTGA AGGGCCGATT CACCATCTCC AGAGACAATT CCAAGAACAC 350
GCTGTATCTG CAAATGAACA GCCTGAGAGC TGAGGACACG GCTGTGTATT 400
ACTGTGCGAG AGATAGGGC TACTACTACA TGGACGTCTG GGGCAAAGGG 450
ACCACGGTCA CCGTCTCCTC AGGTGGAGGC GGTCAGGCG GAGGTGGCTC 500
TGGCGGTGGC GGATCGCAGT CTGTGTTGAC GCAGCCGCC TCAGTGTCTG 550
GGGCCCCAGG ACAGAGGGTC ACCATCTCCT GCACTGGGAG AAGCTCCAAC 600
ATCGGGGCAG GTCATGATGT ACACTGGTAC CAGCAACTTC CAGGAACAGC 650
CCCCAAACTC CTCATCTATG ATGACAGCAA TCGGCCCTCA GGGGTCCCTG 700
ACCGATTCTC TGGCTCCAGG TCTGGCACCT CAGCCTCCCT GGCCATCACT 750
GGGCTCCAGG CTGAAGATGA GGCTGATTAT TACTGCCAGT CCTATGACAG 800
CAGCCTGAGG GGTCGGTAT TCGGCGGAGG GACCAAGGTC ACTGTCTAG 850
GTGCGGCCGC ACATCATCAT CACCATCACG GGGCCGCAGA ACAAAAACTC 900
ATCTCAGAAG AGGATCTGAA TGGGGCCGCA TAG 933

Fig. 15C

100 99 98 97 96 95 94 93 92 91 90 89 88 87 86 85 84 83 82 81 80 79 78 77 76 75 74 73 72 71 70 69 68 67 66 65 64 63 62 61 60 59 58 57 56 55 54 53 52 51 50 49 48 47 46 45 44 43 42 41 40 39 38 37 36 35 34 33 32 31 30 29 28 27 26 25 24 23 22 21 20 19 18 17 16 15 14 13 12 11 10 9 8 7 6 5 4 3 2 1

	signal	Heavy chain
Apo-2.16E2.his	1 MIMITPSFGAFFLEIFNVKKLLFAIPLVVPFYAAQPAMAEVQLVQSGGGV	
Apo-2.20E6.his	1 MIMITPSFGAFFLEIFNVKKLLFAIPLVVPFYAAQPAMAGVQLVESGGGL	
Apo-2.24C4.his	1 MIMITPSFGAFFLEIFNVKKLLFAIPLVVPFYAAQPAMAQVQLVQSGGGV	
		CDR1
		CDR2
Apo-2.16E2.his	51 ERPGGSLRLSCAASGFTFDD <u>DYGM</u> SWRQAPGKGLEW <u>SGINW</u> GGSTGYA	
Apo-2.20E6.his	51 VQPGGSLRLSCAASGFTF <u>SSYWM</u> SWRQAPGKGLEW <u>ANIKODGSEKYYV</u>	
Apo-2.24C4.his	51 VQPGRSLRLSCAASGFI <u>FSSYGMH</u> WVRQAPGKGLEW <u>AGIFYDGGN</u> KYYA	
		CDR3
Apo-2.16E2.his	101 <u>DSVKGRTI</u> SRDNAKNSLYLQMNSLRAEDTAVYYCAK <u>IL</u> ---GAGR ^{WY}	
Apo-2.20E6.his	101 <u>DSVKGRTI</u> SRDNAKNSLYLQMNSLRAEDTAVYYCAR <u>DLLKVKGSSSGW</u>	
Apo-2.24C4.his	101 <u>DSVKGRTI</u> SRDNSKNTLYLQMNSLRAEDTAVYYCARD-----RGYY	
		Light chain
Apo-2.16E2.his	147 <u>F-DLWGKG</u> ITVIVSSGGGGSGGGGGGGGS-SELTQDPAVSVALGQTVRI	
Apo-2.20E6.his	150 <u>F-DPWGRG</u> ITVIVSSGGGGSGGGGGGGGS-SELTQDPAVSVALGQTVRI	
Apo-2.24C4.his	143 <u>YMDVWGKG</u> ITVIVSSGGGGSGGGGGGGSQSVLTOPPSVSGAPGQRVTI	
		CDR1
		CDR2
Apo-2.16E2.his	195 <u>TCQGD</u> SLR---SYYASWYQQKPGQAPVLT <u>Y</u> GKNR <u>PSG</u> IPDRFSGSSSG	
Apo-2.20E6.his	198 <u>TCQGD</u> SLR---SYYASWYQQKPGQAPVLT <u>Y</u> GKNR <u>PSG</u> IPDRFSGSSSG	
Apo-2.24C4.his	193 <u>SCTGRSSNIGAGH</u> DVHWYQQLPGTAPKLL <u>Y</u> DDSNR <u>PSG</u> VPDRFSGSRSG	
		CDR3
Apo-2.16E2.his	242 NTASLTITGAQAEDEADYY <u>CNSRDSSGNHW</u> FGGGT <u>KL</u> TVLGA ^{AAHHHH}	
Apo-2.20E6.his	245 NTASLTITGAQAEDEADYY <u>CNSRDSSGNHW</u> FGGGT <u>KL</u> TVLGA ^{AAHHHH}	
Apo-2.24C4.his	243 TSASLA <u>ITGL</u> QAEDEADYY <u>COSYDSSLRG</u> SVFGGGT <u>KV</u> TVLGA ^{AAHHHH}	
Apo-2.16E2.his	292 HGAAE <u>QKL</u> ISEEDLN <u>GAA</u>	
Apo-2.20E6.his	295 HGAAE <u>QKL</u> ISEEDLN <u>GAA</u>	
Apo-2.24C4.his	293 HGAAE <u>QKL</u> ISEEDLN <u>GAA</u>	

Fig. 16